## In the claims:

For the convenience of the Examiner, all claims being examined, whether or not amended, are presented below.

- 1. (Currently amended) A method of making a cartilaginous prosthesis, comprising seeding a polymeric matrix construct with ehrondrocytes chondrocytes and contacting the seeded construct with hydrophobically modified means for binding patched and thereby activating hedgehog signaling, wherein the seeded construct is contacted with at least 500 ng/ml of said hydrophobically modified means a hedgehog therapeutic.
- 2. (Original) The method of claim 1, wherein the chondrocytes are articular chondrocytes.
- 3. (Currently amended) The method of claim 2, wherein the <u>hydrophobically modified</u> means for binding patched and thereby activating hedgehog signaling comprises hedgehog therapeutic is a naturally-occurring hedgehog polypeptide.
- 4. (**Currently amended**) The method of claim 1/2, wherein the hedgehog therapeutic is a hydrophobically modified hedgehog polypeptide means comprises a dipalmitoyl hedgehog polypeptide.
- 5. (**Currently amended**) The method of claim 1 or 4 3, wherein the hydrophobically modified means hedgehog polypeptide is di-palmitoyl sonic hedgehog or dipalmitoyl indian hedgehog.
- 6. (Currently amended) The method of claim <u>1 or</u> 4, wherein the seeded construct is contacted with dipalmitoyl hedgehog at a concentration of at least 500 ng/ml.
- 7. (Currently amended) The method of claim 6 4, wherein the seeded construct is contacted with dipalmitoyl sonic hedgehog at a concentration of between 500 and 1000 ng/ml.
- 8. (Currently amended) The method of claim 1 or 4, wherein the polymeric matrix is selected from any the group consisting of polyglycolid acid, collagen, dextran sulfate,

polyanhydride, polyorthoester, hyaluronic acid-based polymers, and a glycosaminoglycan.

- 9. (Withdrawn) The method of claim 6, wherein the polymeric matrix is polycolid acid.
- 10. (Withdrawn) A tissue culture system for the production of cartilage comprising a polymeric matrix, chondrocytes adherent to the matrix and a culture medium comprising an amount of a hedgehog therapeutic sufficient to cause an increase in the proteoglycan content of the cartilage, compared to the content in absence of the hedgehog therapeutic.
- 11. (Withdrawn) The tissue culture system of claim 10, in which the hedgehog therapeutic is a naturally occurring hedgehog polypeptide.
- 12. (Withdrawn) The tissue culture system of claim 11, in which the hedgehog polypeptide is sonic hedgehog or indian hedgehog.
- 13. **(Withdrawn)** The tissue culture system of claim 10, in which the hedgehog therapeutic is a hydrophobically modified hedgehog polypeptide.
- 14. (Withdrawn) The tissue culture system of claim 13, in which the hedgehog therapeutic is dipalmitoyl sonic hedgehog or dipalmitoyl indian hedgehog.
- 15. (Withdrawn) The tissue culture system of claim 14, in which the amount of the hedgehog therapeutic is between 500 and 1000 ng/ml.